Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

- (Currently Amended) Method for detecting disease-associated autoantibodies, which are directed at G protein-coupled receptors <u>for diagnosis of autoimmune diseases</u>, <u>comprising</u> characterized in that the method comprises the following steps:
 - a) Bringing bodily fluid into contact with a denaturing agent an agent for precipitating autoantibodies, wherein a fraction of said fluid comprising said autoantibodies is precipitated,
 - b) Bringing the precipitated fraction into contact with a peptide, particularly one comprising Biotin which comprises a sequence or partial sequence of the first and/or second loop of the a G protein-coupled receptor and a tag, whereby a mixture is formed in which the autoantibodies bind said sequence or partial sequence of said peptide,
 - c) Incubating the mixture with a carrier coated with avidin or streptavidin an anti-tag to bind said tag,
 - d) Washing the materials of the carrier,
 - e) Incubating the carrier with anti-lgG antibody subclasses, whereby wherein the anti-lgG antibody is marked for an enzyme reaction or color reaction, and
 - f) Carrying out an <u>said</u> enzyme reaction or color reaction to detect disease-associated autoantibodies, which are directed at said G protein-coupled receptor to diagnose said diseases.
- (Currently Amended) Method of claim 1, characterized in that wherein
 the denaturing agent <u>for precipitating autoantibodies</u> is ammonium sulfate and/or alcohol.
- (Currently Amended) Method of claim 1, characterized in that wherein the carrier is a magnetic particle or an ELISA plate.
- 4. (Currently Amended) Method of claim 1, Appl. No. 10/536,552 Page 4

characterized in that wherein

the autoantibodies are directed against a beta1-adrenergen receptor, a muscarinergen M2 receptor, an angiotensin II AT1 receptor, an alphal-adrenergen receptor, and an endothelin IA receptor, a PAR-1, PAR-2, and/or PAR-3.

5. (Currently Amended) Method of Claims 1 claim 4,

characterized in that wherein

the autoantibodies directed against the beta1-adrenergen receptor are associated with dilatative myocardiopathy, Chagas' myocardiopathy, or myocarditis; the autoantibodies directed against the muscarinergen M2 receptor are associated with dilatative myocardiopathy and/or Chagas' cardiomyopathy; the autoantibodies directed against the angiotensin II AT1 receptor are associated with preeclampsia, and/or malignant hypertension; the autoantibodies directed against the alpha1-adrenergen receptor are associated with essential hypertension, refractory hypertension, pulmonary hypertension and/or psoriasis; and/or the autoantibodies directed against endothelin IA receptor, PAR-1, PAR-2 and/or PAR-3 are associated with Raynaud's syndrome.

6. (Currently Amended) Method of claim 1,

characterized in that wherein

the peptide that comprises a sequence or partial sequence of the first and/or second loop of the receptor is used in the detection of <u>autoantibodies associated with</u> dilatative <u>cardiomyopathy</u> myocardiopathy, myocarditis, essential hypertension, refractory hypertension, pulmonary hypertension, or psoriasis, and that the peptide that comprises a sequence or partial sequence of the second loop of the receptor is used for Chargas' myocardiopathy, dilatative <u>myocardiopathy</u> eardiomyopathy, and/or Raynaud's syndrome.

- 7. (Currently Amended) Method of claim 1,
 - characterized in that wherein
- the autoantibodies associated with dilatative cardiomyopathy are brought into contact with the peptide comprising a sequence or partial sequence of the first and/or second loop of the betal-adrenergen receptor,
- the autoantibodies associated with Chargas' cardiomyopathy are brought into contact with the peptide comprising a sequence or partial sequence of the second loop of the beta1-

Appl. No. 10/536,552

adrenergen receptor,

- the autoantibodies associated with myocarditis are brought into contact with the peptide comprising a sequence or partial sequence of the first and/or second loop of the beta1adrenergen receptor,
- the autoantibodies associated with dilatative cardiomyopathy are brought into contact with the peptide comprising a sequence or partial sequence of the second loop of the muscarinergen M2 receptor,
- the autoantibodies associated with Chargas' cardiomyopathy are brought into contact with the peptide comprising a sequence or partial sequence of the second loop of the muscarinergen M2 receptor,
- the autoantibodies associated with preeclampsia are brought into contact with the peptide comprising a sequence or partial sequence of the second loop of the angiotensin II AT1 receptor,
- the autoantibodies associated with malignant hypertension are brought into contact with the peptide comprising a sequence or partial sequence of the second loop of the angiotensin II AT1 receptor,
- the autoantibodies associated with essential hypertension are brought into contact with the peptide comprising a sequence or partial sequence of the first and/or second loop of the alpha1-adrenergen receptor,
- the autoantibodies associated with refractory hypertension are brought into contact with the peptide comprising a sequence or partial sequence of the first and/or second loop of the alpha1-adrenergen receptor,
- the autoantibodies associated with pulmonary hypertension are brought into contact with the peptide comprising a sequence or partial sequence of the first and/or second loop of the alpha1-adrenergen receptor,
- the autoantibodies associated with psoriasis are brought into contact with the peptide comprising a sequence or partial sequence of the first and/or second loop of the alpha1 -adrenergen receptor,
- the autoantibodies associated with Raynaud's Syndrome are brought into contact with the peptide comprising a sequence or partial sequence of the first and/or second loop of the endothelin IA receptor, PAR-1, PAR-2 and/or PAR-3.

characterized in that

the IgG subclasses are IgG1, IgG2, IgG3 and/or IgG4 subclasses wherein the tag is biotin and the anti-tag is avidin or streptavidin.

- (Currently Amended) Method of claim 1, characterized in that wherein
- in the case of dilatative cardiomyopathy, the IgG3 and/or IgG4 subclasses are used are detected if the peptide comprises a sequence or partial sequence of the first loop, and/or the IgG1 subclass is used is detected if the peptide comprises a sequence or partial sequence of the second loop,
- in the case of Chagas' cardiomyopathy, the IgG1, IgG2, IgG3 and/or IgG4 subclasses are used is detected,
- in the case of myocarditis, the IgG3 and/or IgG4 subclasses are used are detected if the peptide comprises a sequence or partial sequence of the first loop, and/or the IgG1 subclass is used is detected if the peptide comprises a sequence or partial sequence of the second loop,
- in the case of preeclampsia, the IgG3 subclass is used is detected,
- in the case of malignant hypertension, the lgG1 and/or lgG3 subclasses are used are detected,
- in the case of essential hypertension, the IgG1 and/or IgG3 subclasses are used are detected if the peptide comprises a sequence or partial sequence of the first loop, and/or the IgG2 subclass is used is detected if the peptide comprises a sequence or partial sequence of the second loop,
- in the case of refractory hypertension, the IgG1 and/or IgG3 subclasses are used are detected if the peptide comprises a sequence or partial sequence of the first loop, and/or the IgG2 subclass is used is detected if the peptide comprises a sequence or partial sequence of the second loop,
- in the case of pulmonary hypertension, the IgG1, IgG2, IgG3 and/or IgG4 subclasses are used are detected, in the case of psoriasis, the IgG1, IgG2, IgG3 and/or IgG4 subclasses are used are detected, and/or
- in the case of Raynaud's Syndrome, the lgG1 subclass is used is detected.
- 10. (Currently Amended) Method of claim 1,

characterized in that wherein the autoantibodies are concentrated or purified before being Appl. No. 10/536,552

detected-identified.

11. (Currently Amended) Method of claim 10,

characterized in that wherein

the method for concentrating or purifying the autoantibodies comprises the following steps:

- a) i) Obtaining an IgG fraction from bodily fluid,
- b) ii) Bringing the IgG fraction that was obtained into contact with a peptide that comprises a partial sequence of a first or second loop of a G protein-coupled receptor and a tag, whereby a mixture is obtained in which the autoantibody bind said partial sequence of said peptide,
- e) iii) Incubating the mixture with a carrier coated with an anti-tag to bind said tag and that is washed and concentrated, and
- d) iv) Eluting the autoantibodies from the concentrated carrier.
- 12. (Currently Amended) Method of claim 1,

characterized in that wherein

the peptide that comprises the sequence or partial sequence of the first and/or second loop is selected from the group comprising consisting of EYGSFF [SEQ ID NO: 1], SFFCEL [SEQ ID NO: 2], ARRCYND [SEQ ID NO: 3], PKCCDF [SEQ ID NO: 4], AESDE [SEQ ID NO: 5], CYIQFF [SEQ ID NO: 6], EDGECY [SEQ ID NO: 7], VRTVEDGECYIQFFSNAAVTFGTAI [SEQ ID NO: 8], AFHYESQ [SEQ ID NO: 9], ENTNIT [SEQ ID NO: 10], FWAFGR [SEQ ID NO: 11], GRAFCDV [SEQ ID NO: 12], ITEEAGY [SEQ ID NO: 13], ERFCGI [SEQ ID NO: 14], GRIFCD [SEQ ID NO: 15] and/or ITTCHDVL [SEQ ID NO: 16].

- 13. (Cancelled)
- 14. (Cancelled)
- 15. (Currently Amended) Method of claim 1,

characterized in that wherein

the peptide is modified by means of deletion, addition, substitution, translocation, inversion and/or insertion substitution.

Appl. No. 10/536,552 Page 8

- 16. (Withdrawn) Peptide selected from the group comprising EYGSFF, SFFCEL, ARRCYND, PKCCDF, AESDE, CYIQFF, EDGECY, VRTVEDGECYIQFFSNAAVTFGTAI, AFHYESQ, ENTNIT, FWAFGR, GRAFCDV, ITEEAGY, ERFCGI, GRIFCD and/or ITTCHDVL, for use as a medicinal active substance.
- 17. (Withdrawn) Peptide of claim 16,

characterized in that

the peptide is bound by autoantibodies of patients having one of the following diseases: dilatative cardiomyopathy, Chagas' cardiomyopathy, myocarditis, preeclampsia, malignant hypertension, essential hypertension, refractory hypertension, pulmonary hypertension, psoriasis and/or Raynaud's Syndrome.

- (Withdrawn) Peptide of claim 16, characterized in that the peptide is immobilized.
- (Withdrawn) Peptide of claim 16,
 characterized in that
 the peptide the peptide is bound to a solid phase.
- 20. (Withdrawn) Recognition molecule directed against the peptide of claim 16.
- 21. (Withdrawn) Recognition molecule of claim 20, characterized in that it is an antibody, a lectin, an antisense construct, and/or a chelator.
- 22. (Withdrawn) Pharmaceutical composition comprising a peptide selected from the group comprising EYGSFF, SFFCEL, ARRCYND, PKCCDF, AESDE, CYIQFF, EDGECY, VRTVEDGECYIQFFSNAAVTFGTAI, AFHYESQ, ENTNH, FWAFGR, GRAFCDV, ITEEAGY, ERFCGI, GRIFCD and/or ITTCHD and/or a recognition molecule directed against the peptide.
- 23. (Withdrawn) Kit comprising a peptide selected from the group comprising

Appl. No. 10/536,552 Page 9 EYGSFF, SFFCEL, ARRCYND, PKCCDF, AESDE, CYIQFF, EDGECY, VRTVEDGECYIQFFSNAAVTFGTAI, AFHYESQ, ENTNIT, FWAFGR, GRAFCDV, ITEEAGY, ERFCGI, GRIFCD and/or ITTCHD, a recognition molecule directed against the peptide, and/or a pharmaceutical composition comprising the peptide and/or the recognition molecule, if applicable with Instructions for combining the contents of the kit and/or for making available a formulation.

- 24. (Withdrawn) Chromatography device comprising peptides selected from the group comprising EYGSFF, SFFCEL, ARRCYND, PKCCDF, AESDE, CYIQFF, EDGECY, VRTVEDGECYIQFFSNAAVTFGTAI, AFHYESQ, ENTNIT, FWAFGR, GRAFCDV, ITEEAGY, ERFCGI, GRIFCD and/or ITTCHD and/or recognition molecules directed against the peptide.
- 25. (Withdrawn) Device of claim 24, characterized in that the peptides are bound to the solid phase.
- 26. 29. (canceled)
- 30. (Withdrawn) Method for treating an autoimmune disease, selected from the group comprising dilatative cardiomyopathy, Chagas' cardiomyopathy, myocarditis, preeclampsia, malignant hypertension, essential hypertension,refractory hypertension, pulmonary hypertension, psoriasis, Raynaud's syndrome, by means of binding and/or removing antibodies by means of peptidesselected from the group comprising EYGSFF, SFFCEL, ARRCYND, PKCCDF, AESDE, CYIQFF, EDGECY, VRTVEDGECYIQFFSNAAVTFGTAI, AFHYESQ, ENTNIT, FWAFGR, GRAFCDV, ITEEAGY, ERFCGI, GRIFCD and/or ITTCHD, bound to a solid phase.
- (Withdrawn) Method of claim 30, characterized in that

the autoantibodies are directed against betal-adrenergic receptors in the case of dilatative cardiomyopathy, against betal-adrenergic receptors in the case of Chagas' cardiomyopathy, against beta1-adrenergic receptors in the case of myocarditis, against muscarinergic M2 receptors in the case of dilatative cardiomyopathy, against muscarinergic M2 receptors in the case of Chagas' cardiomyopathy, against angiotensin II AT1 receptors in the case of

preeclampsia, against angiotensin II AT1 receptors in the case of malignant hypertension, against alpha1-adrenergic receptors in the case of essential hypertension, against alpha1-adrenergic receptors in the case of refractory hypertension, against alpha1-adrenergic receptors in the case of pulmonary hypertension, against alpha1-adrenergic receptors in the case of psoriasis, and that the autoantibodies are directed against endothelin IA, PAR-1 PAR-2 and/or PAR-3 in the case of Raynaud's Syndrome.

- 32. (Currently Amended) Method for the prophylaxis, diagnosis, therapy, monitoring the progression as well as follow-up treatment of autoimmune diseases selected from the group comprising consisting of dilatative cardiomyopathy, Chagas' cardiomyopathy, myocarditis, preeclampsia, malignant hypertension, essential hypertension, refractory hypertension, pulmonary hypertension, psoriasis and Raynaud's Syndrome, comprising the step of using providing one or more chosen from the following
 - (a) Peptide selected from the group

 comprising consisting of EYGSFF [SEQ ID NO: 1], SFFCEL [SEQ ID NO: 2], ARRCYND

 [SEQ ID NO: 3], PKCCDF [SEQ ID NO: 4], AESDE [SEQ ID NO: 5], CYIQFF [SEQ ID NO:
 6], EDGECY [SEQ ID NO: 7], VRTVEDGECYIQFFSNAAVTFGTAI [SEQ ID NO: 8],

 AFHYESQ [SEQ ID NO: 9], ENTNIT [SEQ ID NO: 10], FWAFGR [SEQ ID NO: 11],

 GRAFCDV [SEQ ID NO: 12], ITEEAGY [SEQ ID NO: 13], ERFCGI [SEQ ID NO: 14],

 GRIFCD [SEQ ID NO: 15] and/or ITTCHDVL [SEQ ID NO: 16],
 - (b) a recognition molecule directed against said peptide,
 - (c) a pharmaceutical composition comprising said peptide and said recognition molecule, or
- (d) a kit comprising one of said peptide, said recognition molecule or said pharmaceutical composition, optionally with instructions for combining the contents of the kit and/or for making available a formulation and
- (e) a chromatrography device comprising said peptide or said recognition molecule, and diagnosing or monitoring the progression of said autoimmune diseases.
- 33. (Withdrawn) Method for the production of a medication for the treatment of autoimmune diseases selected from the group comprising dilatative cardiomyopathy, Chagas' cardiomyopathy, myocarditis, preeclampsia, malignant hypertension, essential hypertension, refractory hypertension, pulmonary hypertension, psoriasis and/or Raynaud's syndrome, comprising the step of using one or more chosen from the following (a) Peptide

selected from the group comprising EYGSFF, SFFCEL, ARRCYND, PKCCDF, AESDE, CYIQFF, EDGECY, VRTVEDGECYIQFFSNAAVTFGTAI, AFHYESQ, ENTNIT, FWAFGR, GRAFCDV, ITEEAGY, ERFCGI, GRIFCD and ITTCHDVL (b) a recognition molecule directed against said peptide (c) a pharmaceutical composition comprising said peptide and said recognition molecule (d)a kit comprising one of said peptide, said recognition molecule or said pharmaceutical composition, optionally with instructions for combining the contents of the kit and/or for making available a formulation and (e) a chromatrography device comprising said peptide or said recognition molecule.

- 34. (Withdrawn) Method for Screening medications, comprising the step of one or more chosen from the following (a) Peptide selected from the group comprising EYGSFF, SFFCEL, ARRCYND, PKCCDF, AESDE, CYIQFF, EDGECY, VRTVEDGECYIQFFSNAAVTFGTAI, AFHYESQ, ENTNIT, FWAFGR, GRAFCDV, ITEEAGY, ERFCGI, GRIFCD and ITTCHDVL (b) a recognition molecule directed against said peptide (c) a pharmaceutical composition comprising said peptide and said recognition molecule (d) a kit comprising one of said peptide, said recognition molecule or said pharmaceutical composition, optionally with instructions for combining the contents of the kit and/or for making available a formulation and (e) a chromatrography device comprising said peptide or said recognition molecule.
- 35. (Currently Amended) Method for detecting, binding, complexing or neutralizing of autoantibodies, directed against beta 1-adrenergen receptor, muscarinergen M2 receptor, angiotensin II AT1 receptor, alpha1-adrenergen receptor, endothelin IA receptor, PAR-1, PAR-2, and/or PAR-3, comprising the step of using

providing a peptide selected from the group comprising EYGSFF [SEQ ID NO: 1], SFFCEL [SEQ ID NO: 2], ARRCYND [SEQ ID NO: 3], PKCCDF [SEQ ID NO: 4], AESDE [SEQ ID NO: 5], CYIQFF [SEQ ID NO: 6], EDGEDY EDGECY [SEQ ID NO: 7], VRTVEDGECYIQFFSNAAVTFGTAI [SEQ ID NO: 8], AFHYESQ [SEQ ID NO: 9], ENTNIT [SEQ ID NO: 10], FWAFGR [SEQ ID NO: 11], GRAFCDV [SEQ ID NO: 12], ITEEAGY [SEQ ID NO: 13], ERFCGI [SEQ ID NO: 14], GRIFCD [SEQ ID NO: 15] and/or ITTCHDVL [SEQ ID NO: 16], and

detecting said autoantibodies.

36. (New) The method of claim 12, wherein the peptides are EYGSFF [SEQ ID NO: 1], *Appl. No.* 10/536,552 Page 12

SFFCEL [SEQ ID NO: 2], ARRCYND [SEQ ID NO: 3] or PKCCDF [SEQ ID NO: 4].

37. (New) The method of claim 1, wherein the peptide comprises a partial sequence of the first and/or second loop of the G protein-coupled receptor and a tag.